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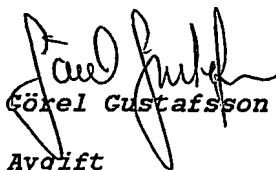
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**Controlled flow assay device and method**

The present invention relates to the field of assay devices or assay systems including components thereof, as well as methods of their use, e.g. a method for detecting an analyte in a blood sample, using such devices or components. The invention in particular relates to such devices and such methods where the flow of liquids is controlled.

**Background**

Analytical and diagnostic determinations are frequently performed on liquid samples, comprising in addition to the analyte of interest, also countless other components, chemical and particulate, which often interfere with the handling of the sample and may influence the quantitative or qualitative determination of the analyte.

For example, numerous clinical diagnostic methods are based on the detection of an analyte in a blood sample. Frequently, such detection is achieved in a disposable assay device, allowing rapid and simple diagnosis. One important application is the wide field of immunology, where analytes are detected with the aid of specific antibodies, capable of binding to the analytes and forming detectable complexes, usually with the aid of ligands aiding the detection.

When performing a test using a blood sample from a patient, many factors need to be considered. Whole blood is prone to clotting, reducing or preventing the desired flow of the sample in the assay device. The red blood cells, even in the absence of clotting, may inhibit or retard flow. Further, red blood cells may inhibit binding between specific binding pair members. Red blood cells also have enzymatic activity, which, depending on the assay employed, may interfere with the signal produced.

Unfortunately, red blood cells present in whole blood also scatter and absorb light thus interfering with assay methodologies which measure either reflected or transmitted light. Other cells may interfere with particular determinations; for example, cholesterol determinations can be effected by cholesterol present in cell membranes.

Consequently many assays involve a step of separating the red blood cells from the plasma, whereupon the assay is carried out on plasma or serum. When the separation is performed before clotting, plasma is obtained. When clotting has occurred before separation, serum is obtained.

- 5 The red blood cells can be separated from plasma through centrifugation, which however requires relatively large volume of sample, and the use of a centrifuge. This is also time consuming and constitutes an additional step of handling the sample, which should be avoided when potentially contagious blood-borne pathogens are involved. Further, the risk of the sample being contaminated by the individuals  
10 handling it, cross-contaminated by parallel sample or mixed up with other samples is increased.

- The most common type of disposable assay device consists of a zone or area for receiving the sample, a reaction zone, and optionally a transport or incubation zone connecting the receiving and reaction zone, respectively. These assay devices are  
15 known as chromatography assay devices or simply referred to as strip tests. They employ a porous material defining a path for fluid flow capable of supporting capillary flow, e.g. a filter material. The sample-receiving zone frequently consists of a more porous material, capable of absorbing the sample, and, when the separation of blood cells is desired, effective to trap the red blood cells. Examples of such materials are  
20 fibrous materials, such as paper, fleece, gel or tissues, comprised e.g. of cellulose, wool, glass fibre, asbestos, synthetic fibres, polymers or mixtures of the same. The transport or incubation zone commonly consists of the same or similar materials, often with another porosity than the sample-receiving zone. Likewise, the reaction zone, which may be integrated with the incubation zone, or constituting the most  
25 distal part thereof, commonly consists of similar, absorbing fibrous materials, or any of the above listed materials.

- In an assay device or strip test, the porous material (-s) is (are) assembled on a carrier, such as a strip of thermoplastic material, paper, cardboard or the like. Further, a cover can be provided, said cover having at least one aperture for  
30 receiving the sample, and an aperture or transparent area for reading the result of the assay.

Nitrocellulose materials are also frequently used as the matrix constituting the transport or reaction zone, connecting the receiving zone and the reaction zone. A significant disadvantage with nitrocellulose is its high non-specific binding of proteins and other bio-molecules. Present test strips however often handle a surplus of sample, reducing the influence of this binding.

#### Prior art

EP 1 371 984 discloses a chromatographic assay device and method for detecting the presence of an analyte in a sample of whole blood, utilizing a red blood cell separating agent to aggregate red blood cells and permit plasma or serum to flow by capillary action. The carrier material is exemplified as a paper (fibrous), or membranes of cellulose, fibreglass, cloth, both naturally occurring and synthetic, as well as porous gels.

Although frequently used and well known in the art, the above carrier materials are associated with many drawbacks. The structure of the materials will always vary between different batches, and also within the material, due to the random distribution of the fibres e.g. in a fibrous material, or cavities e.g. in a gel-like material. Similarly, the chemical properties of the material, e.g. the distribution of chemicals added to the material, will inevitable vary for the same reasons as above.

WO 03/103835 discloses micro fluidic systems comprising a substrate, and, provided on said substrate, at least one flow path interconnecting with functional means in which liquid samples can be subjected to different desired procedures, said flow path comprising a plurality of micro posts protruding from said substrate.

The objective of the present invention is to further develop the micro fluidic systems disclosed in WO 03/103835, and in particular to make available means for controlling or regulating the flow, including enhancing or attenuating the flow of liquid samples, reagents or other components on said substrate.

#### Summary of the invention

The present invention makes available a device for the detection of an analyte in a liquid sample, or a component of such device, said device comprising a flow path

with at least one zone for receiving the sample, and a transport or incubation zone, said zones connected by or comprising an area having projections substantially vertical to its surface, wherein said device further comprises a sink with a capacity of receiving and/or absorbing said liquid sample and supporting or controlling the flow rate of said sample through said transport or incubation zone, said sink comprising an area having projections substantially vertical to its surface, and said sink being adapted to respond to external influence regulating its capacity to receive said liquid sample.

The invention also encompasses embodiments of said device and method, as set forth in the description and claims.

#### Short description of the drawings

The invention will be described in closer detail in the following description, examples, and attached drawings, in which

Fig. 1 schematically shows a cross section of a device according to the invention, where a means for heating causes evaporation of liquid at one end, thus driving the flow along the flow path of the device;

Fig. 2 schematically shows a top view of a device according to the invention, where the heating is performed in zones, starting from the zone most distal to the point where the sample is added;

Fig. 3 schematically shows an embodiment of the invention where two parallel determinations can be performed, the flow rate and thereby incubation time individually adjusted in the two flow paths;

Fig. 4 schematically shows another embodiment where sample, reagents and buffers can be serially added and transported along a flow path in a controlled manner; and

Fig. 5 shows different embodiments, illustrating how the transition from one flow path to another can be arranged: A) using a difference in the geometry of the micro posts; B) using a difference in the height and spacing of the micro posts; and C) using the design of the connection between the flow paths.

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**Description****Definitions**

Before the present device and method is described, it is to be understood that this invention is not limited to the particular configurations, method steps, and materials

5 disclosed herein as such configurations, steps and materials may vary somewhat. It is also to be understood that the terminology employed herein is used for the purpose of describing particular embodiments only and is not intended to be limiting since the scope of the present invention will be limited only by the appended claims and equivalents thereof.

10 It must also be noted that, as used in this specification and the appended claims, the singular forms "a", "an", and "the" include plural referents unless the context clearly dictates otherwise. Thus, for example, reference to a reaction mixture containing "a monoclonal antibody" includes a mixture of two or more antibodies.

15 In describing and claiming the present invention, the following terminology will be used in accordance with the definitions set out herein.

The term "sample" here means a volume of a liquid, solution or suspension, intended to be subjected to qualitative or quantitative determination of any of its properties, such as the presence or absence of a component, the concentration of a component, etc. The sample may be a sample taken from an organism, such as a mammal,

20 preferably a human; or from the biosphere, such as a water sample, or an effluent; or from an technical, chemical or biological process, such as a process of manufacturing, e.g. the production of medicaments, food, feed, or the purification of drinking water or the treatment of waste effluents. The sample may be subjected to qualitative or quantitative determination as such, or after suitable pre-treatment, such

25 as homogenisation, sonication, filtering, sedimentation, centrifugation, heat-treatment etc. Typical samples in the context of the present invention are body fluids such as blood, plasma, serum, lymph, urine, saliva, semen, gastric fluid, sputum, tears etc.; environmental fluids such as surface water, ground water, sludge etc.; and process fluids such as milk, whey, broth, nutrient solutions, cell culture medium, etc. The

30 present invention is applicable to all samples, but preferably to samples of body fluids, and most preferably to whole blood samples.

The determination may be for any purpose, such as diagnostic, environmental, quality control, regulatory, or research purposes. The invention can be applied to competitive (inhibition) as well as non-competitive (non-inhibition) test variants.

Examples of diagnostic determinations include, but are not limited to, the

- 5 determination of biospecific affinity reactants such as a cell or virus or a part thereof. In particular antigens, such as an immunoglobulin (Ig) or an antibody may be mentioned. For immunoglobulins the determination may relate to a certain Ig and/or certain Ig subclass.

- 10 The term "analyte" means any substance that is measured quantitatively or qualitatively. The invention in particular relates to analytes exhibiting biospecific affinity, such as individual members on reactant pairs: antigen/hapten - antibody; biotin - avidin/streptavidin; two complimentary nucleic acid chains etc.

- 15 The terms "zone", "area" and "site" are used in the context of this description, examples and claims to define parts of the flow path on a substrate, either in prior art devices or in a device according to the invention.

- 20 The term "reaction" is used to define any reaction taking place between components of a sample and at least one reagent or reagents on or in said substrate, or between two or more components present in said sample. The term "reaction" is in particular used to define the reaction, taking place between an analyte and a reagent as part of the qualitative or quantitative determination of said analyte.

The term "substrate" here means the carrier or matrix to which a sample is added, and on or in which the determination is performed, or where the reaction between analyte and reagent takes place.

- 25 The term "chemical functionality" comprises all chemical reactions between components in solution, between components in solution and components in solid phase, immobilised directly to the substrate, or immobilised to particles, reactions between components in a sample, reagent or buffer, with each other, the ambient or the like, taking part in the pre-treatment, incubation or detection of an analyte in a sample.

- 30 The term "biological functionality" comprises all biological interactions between a component in a sample and a reagent on the substrate, such as catalysis, binding, internalisation, activation etc.



The term "physical functionality" here comprises functionalities involved in reactions and interactions other than those that are mainly chemical or biological. Examples include diameter, height, shape, cross section, surface topography and surface patterns, the number of projections per unit area, wetting behavior of the surface of said projections, or a combination thereof, and/or other functionalities influencing the flow, retention, adhesion or rejection of components of the sample.

The distinctions between chemical, biological and physical interactions are not always clear, and it is possible that an interaction - such as an interaction between a component in a sample and a reagent on the substrate - involves chemical, biological as well as physical elements.

The terms hydrophilic and hydrophobic, as in hydrophilic or hydrophobic compounds, hydrophilic or hydrophobic interactions etc., have the meaning generally understood by a person skilled in the art, and corresponding to that used in generally recognised textbooks.

#### Description of preferred embodiments

The present invention makes available a device for handling liquid samples, said device comprising a flow path with at least one zone for receiving the sample, and a transport or incubation zone, said zones connected by or comprising an area having projections substantially vertical to its surface, wherein said device further comprises a sink with a capacity of receiving said liquid sample and supporting or controlling the flow rate of said sample through said transport or incubation zone, said sink comprising an area having projections substantially vertical to its surface, and said sink being adapted to respond to external influence regulating its capacity to receive said liquid sample.

The device according to the invention can also comprise two or more flow paths, each connected to a sink, said device being adapted for performing multiple analyses on one sample. In this case, each flow path comprises a reaction zone, and individual reagents, such as conjugates, buffers etc are added to or stored in each flow path or reaction zone. A device according to this embodiment is advantageous, as it makes it possible to perform multiple analyses in parallel or substantially in parallel, starting from one sample, added to one device.

According to one embodiment, multiple reagents, buffers, etc are serially added to one flow path. This means that each one of sample, reagent, buffer etc will travel along the flow path and pass the reaction zone in a predetermined order. Using the sink, and in particular a heated sink, the flow speed of each component can be controlled. This makes it possible to perform an e.g. analysis involving a slow pre-treatment, an incubation of a predetermined length, followed by a rapid rinse etc, only to mention an example.

According to the invention, the external influence regulating the capacity of said sink to receive said liquid sample is chosen among heating, cooling, irradiation with visible light, infrared irradiation, vibration, and the application of an electric current.

According to an embodiment of the invention, the sink is divided into sub sections, suitable for being serially subjected to said external influence. This is advantageous e.g. in instances where the sample tends to coagulate, denaturate or simply to dry in the sink during the heating. By serially heating sections of the sink, starting from the most distal one, it is possible to retain the aspirating or absorbing capacity of the sink.

One embodiment of the invention concerns the provision of means or design measures creating a preferred direction of flow within the device. Such means or measures can comprise vertical projections have different cross section in different zones of the flow path, different spacing between said projections, different chemical or biochemical treatment of said projections, a difference in the level of said flow paths, such as forming steps or thresholds between different sections of the flow path etc.

The direction of flow, e.g. the prevention of undesired back flow of sample is chosen among suitable design of the flow paths, the cross section of the substantially vertical projections, an external influence chosen among heating, cooling, irradiation with visible light, infra red irradiation, vibration, and the application of an electric current, or a combination thereof, acting on at least part of said flow paths.

The present invention makes available a chemical or biochemical assay involving a reaction between an analyte in a sample and one or more reagents, wherein the sample is added to a device as defined above. The reaction between said analyte and one or more reagent may be any conventional reaction, presently performed on

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a solid substrate or in a carrier material. This also comprises assays where a device according to the invention is used for the pre-treatment of the sample.

The present invention also makes available chemical or biochemical assay involving a reaction between an analyte in a sample and one or more reagents, wherein said  
5 reaction itself and/or the reading of the result is performed on a device as defined above.

The invention also makes available a method for handling liquid samples, wherein a device as defined above is used.

The device according to the invention is advantageously used in analytical  
10 applications where the liquid sample contains particulate matter, such as cells, tissue debris, organic or inorganic matter, other contamination etc, which is desired to separate from the bulk of the sample. One important application is when the liquid sample is whole blood and in such cases, the lateral capillary flow involves the separation of red blood cells from plasma without significant rupture of said cells.  
15 According to one embodiment, such separation in general, and in particular the gentle separation of red blood cells, is achieved in a gradient of projections wherein the spacing decreases from about 7  $\mu\text{m}$  to about 1  $\mu\text{m}$  over the length of said filtering zone.

According to one embodiment said receiving zone forms a basin for components  
20 separated from the lateral flow, e.g. particulate matter or cells prevented from passing between the projections, or entering that space only to a limited degree.

According to another embodiment the particulate matter travels with the lateral flow. In applications where said liquid sample is whole blood, it is important that said lateral capillary flow involves the transportation of red blood cells without significant  
25 rupture of said cells. This is achieved by the present invention through the control of one or more of the parameters of the projections, such as the height, diameter and reciprocal spacing, as well as the chemical or biochemical derivatisation of the projections.

The spacing between said projections can be varied depending on the intended use  
30 and the properties of the liquid sample, as well as the properties of components to be separated or transported, and preferably is in the interval of 1 to 100  $\mu\text{m}$ , more preferably in the interval of 1 to 50  $\mu\text{m}$ . The distance between said projections can be

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chosen by a skilled person, considering which sample the device is intended for, the properties of said sample, and the properties of the components that are to be separated.

The device according to the present invention is built on a plastic substrate, preferably thermoplastic, or a substrate having a plastic upper layer. This can in turn be coated or derivatised, e.g. using techniques such as sputtering, vapour deposition and the like, and given a coating of silicon, a metal or other. The present invention can also be made of silicon substrates. According to a preferred embodiment the substrate is given a hydrophilic treatment or coating, e.g. by subjecting the substrate to an oxidative treatment, such as e.g. gas plasma treatment, coating with a hydrophilic substance such as silicon oxide, hydrophilic polymers such as dextran, polyethylene glycol, heparin and derivatives thereof, detergents, biologic substances such as polymers, etc.

Consequently, according to one embodiment of the invention, said projections or at least a sub-set thereof are provided with a chemical, biologic or physical functionality. The projections may have chemically reactive groups on their surface. The projections may also have substances with biological affinity bound to their surface.

According to another embodiment, the projections carry structures or groups chosen among hydrophilic groups, hydrophobic groups, positively and/or negatively charged groups, silicon oxide, carbohydrates, amino acids, nucleic acids, and macromolecules, or combinations thereof.

According to yet another embodiment, the projections have a physical property selected from the projection diameter, height, reciprocal spacing, shape, cross section, surface coating, the number of projections per unit area, wetting behavior of the surface of said projections, or a combination thereof, according to the desired end use of the substrate.

According to another embodiment, particles are provided chemically or physically bound to the substrate, or mechanically trapped within a region comprising a plurality of projections. Said particles are chosen among commercially available particles, so called beads, and may have a core of glass, metal or polymer, or a combination of these, and they optionally carry on their surface chemical or biological moieties,

such as polyclonal antibodies, monoclonal antibodies, amino acids, nucleic acids, carbohydrates or combinations thereof.

The present invention also makes available a device suitable for use in or together with a device for detection of an analyte in a liquid sample, wherein said device has  
5 projections substantially vertical to its surface, said projections having a height, diameter and reciprocal spacing such, that said device is capable of separating components of said liquid sample while achieving a lateral flow of said liquid sample. This device may have one or more of the properties and functionalities described above, depending on its intended use.

10 This device may be used separately, in association with, or integrated in a device for the analysis of a liquid sample. This device may function as a pre-treatment step in or before a conventional analysis.

The present invention also makes available a method for performing an assay on a liquid sample, said sample being applied to a substrate having a zone for receiving  
15 the sample, which is in fluid connection with a reaction zone, and optionally a transport or incubation zone connecting the receiving and reaction zone, respectively, wherein said substrate is a non-porous substrate, and said receiving zone, reaction zone and optional transport or incubation zone consist of areas of projections substantially vertical to said surface, and having a height, diameter, and  
20 reciprocal spacing such, that lateral capillary flow of said liquid sample in said zone is achieved.

According to one embodiment of this method, a filtering step is performed following the addition of the sample, said filtering effected in a filtering zone by projections substantially vertical to the surface of said substrate, the projections having a height,  
25 diameter, and reciprocal spacing forming a gradient with regard to the diameter, and/or reciprocal spacing such that components of the sample are gradually retained.

This method can be used for all applications where components of a liquid sample need to be separated from the bulk of the sample. The method is however particularly suitable for applications where said liquid sample is whole blood and said  
30 lateral capillary flow involves the separation of red blood cells from plasma without significant rupture of said cells.

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One way to achieve a gentle separation of components of the sample, is to subject the sample to a filtering zone where the spacing between the projections gradually decreases. In applications where the lateral capillary flow involves the separation of red blood cells from plasma, it is important that this takes place without significant rupture of said red blood cells. To achieve this, in applications involving the separation of red blood cells, the spacing preferably decreases from about 7  $\mu\text{m}$  to about 1  $\mu\text{m}$  over the length of said filtering zone.

According to one embodiment of the method, components separated from the lateral flow are retained in a basin, substantially prevented from entering the filtering zone.

Another embodiment, in applications where said liquid sample is whole blood, is a method of achieving a lateral capillary flow involving the transportation of red blood cells without significant rupture of said cells. And if needed a flow of buffer liquid can be used to reduce the amount of cells in the reaction zone, promoting the detection step.

The invention also makes available a method for performing an assay on a liquid sample, in particular a sample of whole blood, wherein said sample is added to a device as described above.

The device according to the present invention surprisingly replaces prior art devices where the substrate, and/or one or more of said zones were made of a porous material such as nitrocellulose, cellulose, asbestos fibres, glass fibres and the like.

A general embodiment is schematically shown in Fig. 1, showing a part of a device where the surface of a substrate 2 has a flow path 4 comprising projections substantially vertical to said surface, and having a height, diameter and reciprocal spacing such, that lateral capillary flow is achieved. A reaction zone 6 is provided on said surface, and it is desired that sample and optionally reagents, buffers is/are transported laterally so that they pass said reaction zone. It is further desired that the rate, at which said reagents etc pass the reaction zone is controlled. To this end, a sink 8 is provided at the distal end of the substrate, in relation to the place where sample is added. This sink is then heated by heating means 10, in order to evaporate liquid and control or increase the flow rate of the sample.

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Another embodiment is illustrated in Fig. 2, showing a part of a substrate 2 having a flow path 4 and a reaction zone 6. The sink is however divided into zones 12, 14, 16, and 18, which can be heated consecutively, starting from the most distal zone 12.

Yet another embodiment is illustrated in Fig. 3, where sample is added to one location 20 in fluid connection with two flow paths 4' and 4'', each flow path passing a reaction zone 6' and 6'', and ending in a sink, 22' and 22'', respectively. By giving the sinks 22' and 22'' different capacity, or by subjecting them to heating, it becomes possible to achieve different flow rates in the flow paths 4' and 4''. This is advantageous in applications where two or more determinations are to be performed on the same sample, and each determination has its own requirements as to incubation time, reaction time, flow rate etc. While Fig. 3 only shows two flow paths, it is understood that three, four or several parallel flow paths could be provided.

Fig. 4 illustrates an embodiment where sample is added to one location 24, in a system 28 forming a fluid connection between sample addition 24, a flow path 4, a reaction zone 6 and a sink 26, where a first wash buffer  $W_1$  stored in or added to another location 30, a conjugate C stored in or added to 32, and a second wash buffer  $W_2$  stored in or added to 34, can be serially introduced onto said flow path 4. When  $W_1$ ,  $W_2$ , and C are added simultaneously, the distance between 30, 32, and 34; and the flow path 4, respectively, can be varied in order to influence the time it takes  $W_1$ ,  $W_2$ , and C to reach the flow path. When  $W_1$ ,  $W_2$ , and C are stored on the surface, at the locations 30, 32, and 34, respectively, the release of these components can be controlled by the provision of meltable seals, the heating or cooling of the components, or other means, influencing the time it takes  $W_1$ ,  $W_2$ , and C to reach the flow path. While 30, 32, and 34 are schematically shown in Fig. 4 to be at an approximately equal distance from the flow path 4, and approximately vertical thereto, it is contemplated that they can be at different distances, at an angle to the flow path, or even partially parallel thereto. Centrifugation is contemplated as one means to deliver one or all of  $W_1$ ,  $W_2$ , and C to the flow path.

Fig. 5A schematically illustrates an embodiment where the transition from one flow path 36 into another flow path 38 is characterised in that the substantially vertical projections in said flow paths have different properties, said properties chosen so that the direction of flow is controlled, e.g. creating a preferred direction of flow. Different

properties in this respect can be different cross section, height, orientation, spacing, and surface chemistry of the projections, or a combination thereof.

Fig. 5B schematically illustrates another embodiment where not only the properties of said projections, but also the level of the flow paths are different, again in order to  
5 create a preferred direction of flow.

Finally, Fig. 5C schematically illustrates an embodiment where the geometry of the respective flow paths 44 and 46 has been designed to create a preferred direction of flow.

The above embodiments may also be combined, e.g. by taking features from one  
10 embodiment and introducing these in another.

#### Advantages of the invention

An advantage of the device according to the invention is the possibility to accurately control the flow rate, including possibilities to stop and start the flow. This in turn makes it possible to perform simultaneous or sequential reactions, either in series or  
15 in parallel, in a controlled fashion.

Another advantage of the device is that, due to the open, regular structure and the defined properties of the capillary flow zones, the addition of reagents these zones or the derivatisation of the surface of the projections is greatly simplified.

Yet another advantage of the device is the uniformity of the structure not only within a  
20 single device, but also between all devices produced. This result in significantly increased reliability and repeatability of the assays built on the inventive device.

An important advantage of the inventive device is that the degree of separation, from none to total, of the blood cells, can be accurately controlled.

The inventive device has many advantages with respect to the manufacturing  
25 process. All capillary zones can be made in one step and no assembly of parts is required. Optionally, a cover having at least one aperture for sample addition and one reading the result of the assay can be placed over the substrate and the capillary zones.

Although the invention has been described with regard to its preferred embodiments,  
30 which constitute the best mode presently known to the inventors, it should be understood that various changes and modifications as would be obvious to one



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having the ordinary skill in this art may be made without departing from the scope of the invention which is set forth in the claims appended hereto.

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**Claims**

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1. A device for handling liquid samples, said device comprising a flow path with at least one zone for receiving the sample, and a transport or incubation zone, said zones connected by or comprising an area having projections substantially vertical to its surface, characterized in that said device further comprises a sink with a capacity of receiving said liquid sample and supporting or controlling the flow rate of said sample through said transport or incubation zone, said sink comprising an area having projections substantially vertical to its surface, and said sink being adapted to respond to external influence regulating its capacity to receive said liquid sample.
2. The device according to claim 1, wherein two or more flow paths are provided, each connected to a sink, said device adapted for performing multiple analyses on one sample.
3. The device according to claim 2, wherein said multiple analyses are performed in parallel.
4. The device according to claim 1, wherein multiple reagents, buffers, etc can be serially added to one flow path.
5. The device according to any one of claims 1 — 4, wherein the external influence regulating the capacity of said sink to receive said liquid sample is chosen among heating, cooling, irradiation with visible light, infra red irradiation, vibration, and the application of an electric current.
6. The device according to claim 5, wherein the sink is divided into sub sections, suitable for being serially subjected to said external influence.
7. The device according to claim 5 or 6, wherein the sink is heated to evaporate liquid sample therefrom.
8. The device according to any one of claims 1 — 7, wherein the substantially vertical projections have different cross section in different zones of the flow path.
9. The device according to claim 4, wherein the back flow of sample is prevented by suitable design of the flow paths, the cross section of the substantially vertical projections, an external influence chosen among heating, cooling, irradiation with visible light, infra red irradiation, vibration, and the application of an electric current, or a combination thereof, acting on at least part of said flow paths.

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10. A chemical or biochemical assay involving a reaction between an analyte in a sample and one or more reagents, characterized in that the sample is added to a device according to any one of claims 1 - 9.
11. A chemical or biochemical assay involving a reaction between an analyte in a sample and one or more reagents, characterized in that said reaction is performed on a device according to any one of claims 1 - 9.
12. A method for handling liquid samples, characterized in that a device according to any one of claims 1 - 9 is used.

**Abstract**

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A device for handling liquid samples, comprising a flow path with at least one zone for receiving the sample, and a transport or incubation zone, said zones connected by or comprising an area having projections substantially vertical to its surface, is provided with a sink with a capacity of receiving said liquid sample, said sink comprising an area having projections substantially vertical to its surface, and said sink being adapted to respond to an external influence regulating its capacity to receive said liquid sample.

(Fig. 1)

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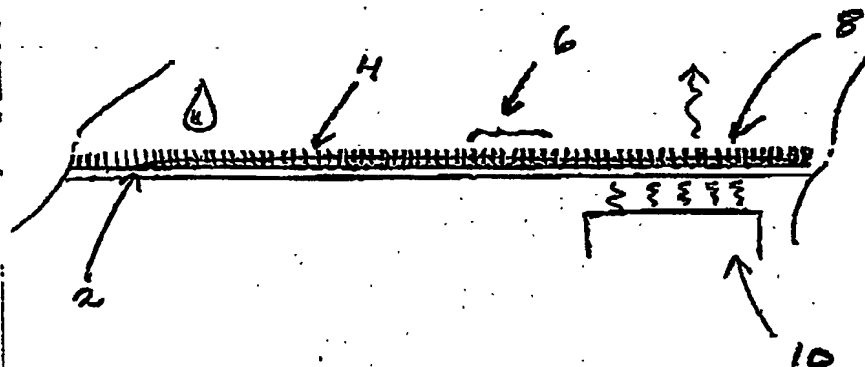


Fig 1

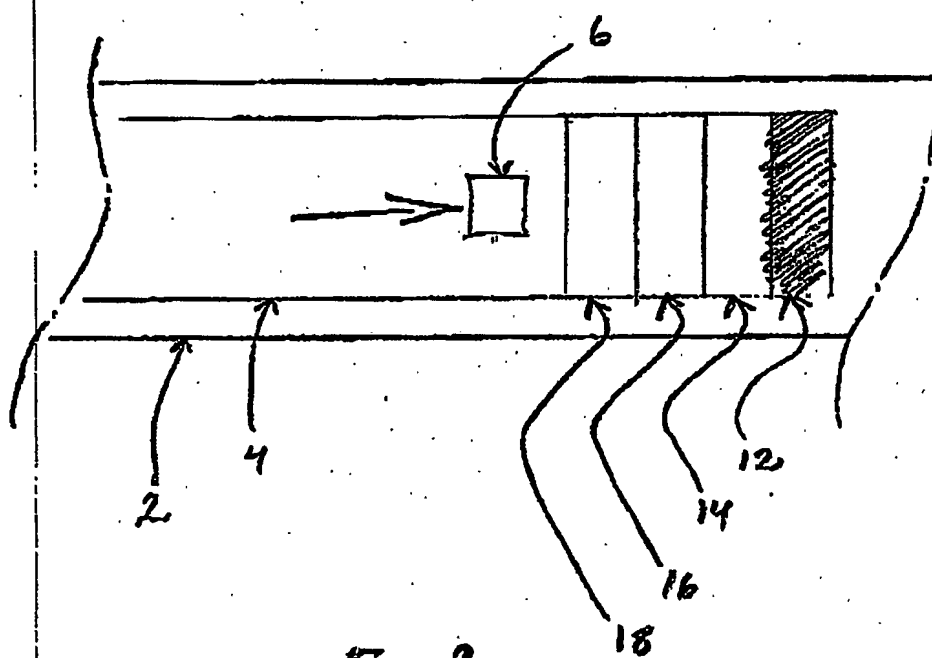


Fig 2

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## Huvudoxen Kassen

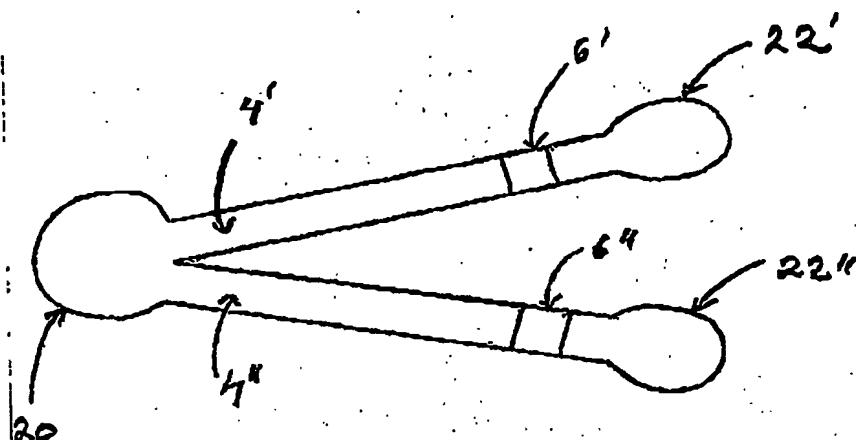


Fig 3

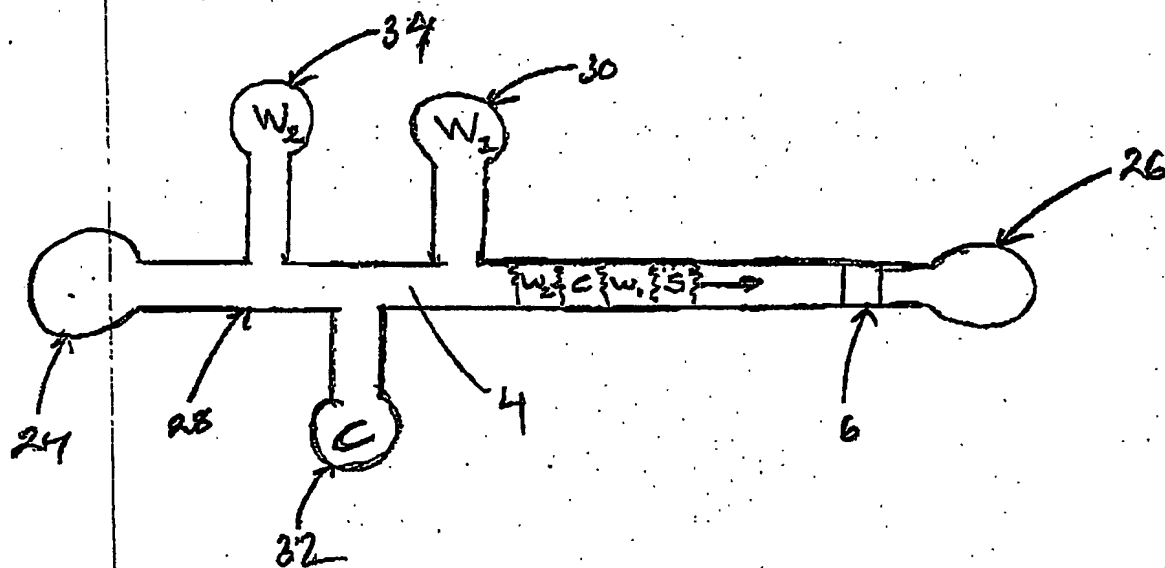


Fig 4

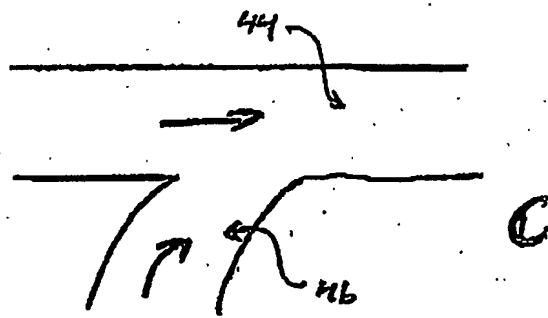
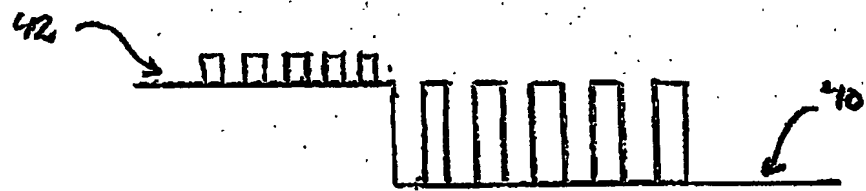
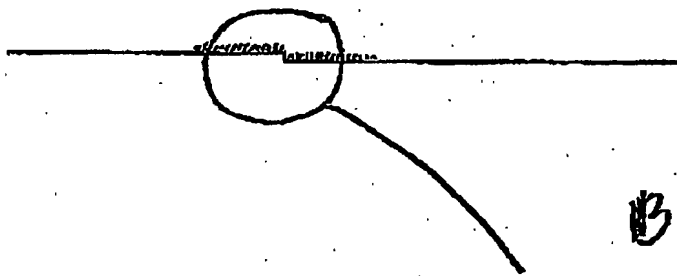
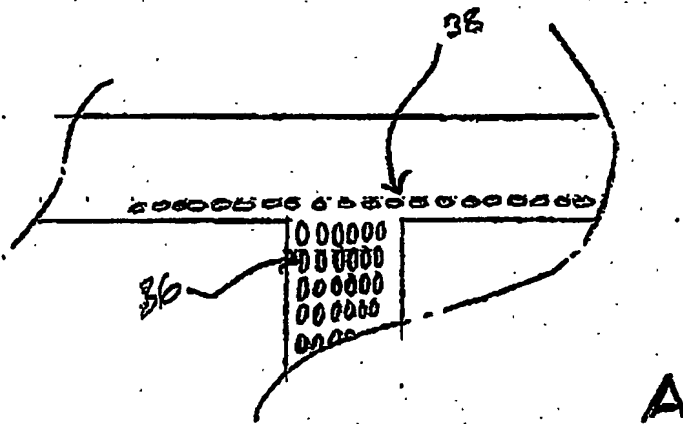


Fig 5

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